



SEQUENCE LISTING

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Anderson, Wesley
Ansari, Aslam

<120> METHODS OF ANALGESIA

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<160> 53

<170> PatentIn version 3.1

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<220>

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<222> (6)..(6)

<223> Polymer connected to epsilon-amino group

<400> 1

Tyr Gly Gly Phe Met Lys

1

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<220>

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<222> (1)..(1)

<223> Polymer connected to alpha-amino group

<220>

<221> MOD_RES

<222> (6)..(6)

<223> Polymer connected to epsilon-amino group

<400> 2

Tyr Gly Gly Phe Met Lys
1 5

<210> 3

<211> 6

<212> PRT

<213> Artificial sequence

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<220>

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<222> (1)..(1)

<223> Polymer connected to alpha-amino group

<400> 3

Tyr Gly Gly Phe Met Lys
1 5

<210> 4

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<222> (1)..(1)

<223> ACETYLATION

<220>

<221> MOD_RES

<222> (6)..(6)

<223> AMIDATION

<400> 4

Phe Arg Trp Trp Tyr Lys
1 5

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<223> ACETYLATION

<220>
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<222> (6)..(6)
<223> AMIDATION

<400> 5

Arg Trp Ile Gly Trp Lys
1 5

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<220>
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<222> (6)..(6)
<223> AMIDATION

<220>
<221> MISC_FEATURE
<222> (6)..(6)
<223> Xaa can be any of the twenty naturally occurring amino acids

<400> 6

Trp Trp Pro Lys His Xaa
1 5

<210> 7
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<222> (4)..(4)
<223> AMIDATION

<220>
<221> MISC_FEATURE
<222> (4)..(4)

<223> Xaa is either Lys or Arg

<400> 7

Trp Trp Pro Xaa

1

<210> 8

<211> 6

<212> PRT

<213> Artificial sequence

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<220>

<221> MOD_RES

<222> (6)..(6)

<223> AMIDATION

<220>

<221> MISC_FEATURE

<222> (6)..(6)

<223> Xaa can be any one of the naturally occurring amino acids

<400> 8

Tyr Pro Phe Gly Phe Xaa

1

5

<210> 9

<211> 7

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<220>

<221> MOD_RES

<222> (1)..(5)

<223> Amino acids are in the D-form

<220>

<221> MOD_RES

<222> (6)..(6)

<223> n is 0 or 1

<220>

<221> MISC_FEATURE

<222> (7)..(7)

<223> Xaa is Gly or the D-form of a naturally occurring amino acid

<220>
<221> MOD_RES
<222> (7)..(7)
<223> Amidation

<400> 9

Ile Met Ser Trp Trp Gly Xaa
1 5

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<222> (1)..(4)
<223> Amino acids are in the D-form

<220>
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<222> (6)..(6)
<223> Xaa is Gly or the D-form of a naturally occurring amino acid

<220>
<221> MOD_RES
<222> (6)..(6)
<223> AMIDATION

<400> 10

Ile Met Thr Trp Gly Xaa
1 5

<210> 11
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<220>
<223> Synthetic construct

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<222> (2)..(2)
<223> Xaa is A1, wherein A1 is the D-form of Nve or Mle

<220>
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<222> (3)..(3)

<223> Xaa is B2, wherein B2 is Gly, Phe, or Trp

<220>

<221> MISC_FEATURE

<222> (4)..(4)

<223> Xaa is C3, wherein C3 is Trp or Nap

<220>

<221> MOD_RES

<222> (4)..(4)

<223> AMIDATION

<400> 11

Tyr Xaa Xaa Xaa

1

<210> 12

<211> 3

<212> PRT

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<223> Synthetic construct

<220>

<221> MOD_RES

<222> (1)..(1)

<223> Tyr has at its N-terminus a Me-x-H-y-N group, wherein x is 0, 1, or 2; and y is 0, 1, or 2, with the proviso that x and y is never greater than 2

<220>

<221> MOD_RES

<222> (1)..(2)

<223> The amine between the first Tyr and the second Tyr is methylated, wherein z is 0 or 1

<220>

<221> MISC_FEATURE

<222> (3)..(3)

<223> Xaa is Phe or (D) Phe

<220>

<221> MOD_RES

<222> (3)..(3)

<223> AMIDATION

<400> 12

Tyr Tyr Xaa

1

<210> 13
<211> 6
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<213> Artificial sequence

<220>
<223> Synthetic construct

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<222> (4)..(4)
<223> Xaa is D4, wherein D4 is Lys or Arg

<220>
<221> MOD_RES
<222> (5)..(5)
<223> His is His-z, wherein z is 0 or 1

<220>
<221> MISC_FEATURE
<222> (6)..(6)
<223> Xaa is Xaa-z, wherein Xaa is a naturally occurring amino acid and
z is 0 or 1

<220>
<221> MOD_RES
<222> (6)..(6)
<223> AMIDATION

<400> 13

Trp Trp Pro Xaa His Xaa
1 5

<210> 14
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<220>
<223> Synthetic construct

<220>
<221> MISC_FEATURE
<222> (2)..(2)
<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<400> 14

Tyr Xaa Phe Phe
1

<210> 15

<211> 4
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<220>
<223> Synthetic construct

<220>
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<222> (2)..(2)
<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
<221> MOD_RES
<222> (4)..(4)
<223> AMIDATION

<400> 15

Tyr Xaa Phe Phe
1

<210> 16
<211> 4
<212> PRT
<213> Artificial sequence

<220>
<223> Synthetic construct

<220>
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<222> (1)..(1)
<223> Tyr is Tyr(N-alpha-Me), i.e. N-alpha-methyl tyrosine

<220>
<221> MISC_FEATURE
<222> (2)..(2)
<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<400> 16

Tyr Xaa Phe Phe
1

<210> 17
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<220>
<223> Synthetic construct

<220>
<221> MOD_RES

<222> (1)..(1)
 <223> Tyr is Tyr(N-alpha-Cmp), i.e. N-alpha-cyclopropylmethyltyrosine

<220>
 <221> MISC_FEATURE
 <222> (2)..(2)
 <223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<400> 17

Tyr Xaa Phe Phe
 1

<210> 18
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<220>
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 <222> (1)..(1)
 <223> Tyr is Tyr(N-alpha-hex), i.e. N-alpha-hexyltyrosine

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 <222> (2)..(2)
 <223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<400> 18

Tyr Xaa Phe Phe
 1

<210> 19
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 <222> (1)..(1)
 <223> Tyr is Tyr(N-alpha-Et2), i.e. N-alpha-diethyltyrosine

<220>
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 <222> (2)..(2)
 <223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<400> 19

Tyr Xaa Phe Phe
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<210> 20

<211> 4

<212> PRT

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<223> Synthetic construct

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<222> (1)..(1)

<223> Tyr is Dmt, i.e. 2,6-dimethyltyrosine

<220>

<221> MISC_FEATURE

<222> (2)..(2)

<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

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Tyr Xaa Phe Phe
1

<210> 21

<211> 4

<212> PRT

<213> Artificial sequence

<220>

<223> Synthetic construct

<220>

<221> MOD_RES

<222> (1)..(1)

<223> Tyr is Dmt, i.e. 2,6-dimethyltyrosine

<220>

<221> MISC_FEATURE

<222> (2)..(2)

<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>

<221> MOD_RES

<222> (4)..(4)

<223> AMIDATION

<400> 21

Tyr Xaa Phe Phe

1

<210> 22

<211> 4

<212> PRT

<213> Artificial sequence

<220>

<223> Synthetic construct

<220>

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<222> (1)..(1)

<223> Tyr is H-Tyr(3-F), i.e. 3-fluorotyrosine

<220>

<221> MISC_FEATURE

<222> (2)..(2)

<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<400> 22

Tyr Xaa Phe Phe

1

<210> 23

<211> 4

<212> PRT

<213> Artificial sequence

<220>

<223> Synthetic construct

<220>

<221> MOD_RES

<222> (1)..(1)

<223> Tyr is H-Tyr(3-Cl), i.e. 3-chlorotyrosine

<220>

<221> MISC_FEATURE

<222> (2)..(2)

<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<400> 23

Tyr Xaa Phe Phe

1

<210> 24

<211> 4

<212> PRT

<213> Artificial sequence

<220>
 <223> Synthetic construct

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 <222> (1)..(1)
 <223> Tyr is H-Tyr (3-Br), i.e. 3-bromotyrosine

 <220>
 <221> MISC_FEATURE
 <222> (2)..(2)
 <223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<400> 24

Tyr Xaa Phe Phe
 1

<210> 25
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<220>
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 <222> (1)..(1)
 <223> Tyr is Dmt, i.e. 2,6-dimethyltyrosine

<220>
 <221> MISC_FEATURE
 <222> (2)..(2)
 <223> Xaa is Tic-psi-[CH2-], i.e. 3-methyl-1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
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 <222> (2)..(3)
 <223> nonpetidyl bond

<400> 25

Tyr Xaa Phe Phe
 1

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<222> (1)..(1)
<223> Tyr is Dmt, i.e. 2,6-dimethyltyrosine

<220>
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<222> (2)..(2)
<223> Xaa is Tic-psi-[CH2-], i.e. 3-methyl-1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
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<222> (2)..(3)
<223> nonpeptidyl bond

<220>
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<222> (4)..(4)
<223> AMIDATION

<400> 26

Tyr Xaa Phe Phe
1

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<220>
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<222> (2)..(2)
<223> Xaa is Tic-psi-[CH2-], i.e. 3-methyl-1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
<221> MOD_RES
<222> (3)..(3)
<223> Phe is -NCH3]Phe, i.e. N-methylphenylalanine

<400> 27

Tyr Xaa Phe Phe
1

<210> 28
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 <220>
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 <220>
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 <222> (2)..(2)
 <223> Xaa is Tic-psi-[CH2-], i.e. 3-methyl-1,2,3,4-tetrahydroisoquinoli
 ne-3-carboxylic acid

<220>
 <221> MOD_RES
 <222> (3)..(3)
 <223> Phe is -NH]Hfe, i.e. homophenylalanine

<400> 28

Tyr Xaa Phe Phe
 1

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<220>
 <223> Synthetic construct

<220>
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 <222> (1)..(1)
 <223> Tyr is Tyr(NMe), i.e. N-methyltyrosine

<220>
 <221> MISC_FEATURE
 <222> (2)..(2)
 <223> Xaa is Tic-psi-[CH2-], i.e. 3-methyl-1,2,3,4-tetrahydroisoquinoli
 ne-3-carboxylic acid

<220>
 <221> MOD_RES
 <222> (3)..(3)
 <223> Phe is -NH]Hfe, i.e. homophenylalanine

<400> 29

Tyr Xaa Phe Phe
 1

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<223> Synthetic construct

<220>

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<222> (2)..(2)

<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>

<221> MOD_RES

<222> (3)..(3)

<223> Gly is Phg, i.e. phenylglycine

<400> 30

Tyr Xaa Gly Phe

1

<210> 31

<211> 4

<212> PRT

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<220>

<223> Synthetic construct

<220>

<221> MISC_FEATURE

<222> (2)..(2)

<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

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Tyr Xaa Trp Phe

1

<210> 32

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<220>

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<222> (2)..(2)

<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>

<221> MOD_RES

<222> (4)..(4)

<223> AMIDATION

<400> 32

Tyr Xaa Trp Phe

1

<210> 33

<211> 4

<212> PRT

<213> Artificial sequence

<220>

<223> Synthetic construct

<220>

<221> MISC_FEATURE

<222> (2)..(2)

<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<400> 33

Tyr Xaa His Phe

1

<210> 34

<211> 4

<212> PRT

<213> Artificial sequence

<220>

<223> Synthetic construct

<220>

<221> MISC_FEATURE

<222> (2)..(2)

<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>

<221> MOD_RES

<222> (3)..(3)

<223> Ala is 2-Nal, i.e. 3-(2'-naphthyl)alanine

<400> 34

Tyr Xaa Ala Phe

1

<210> 35

<211> 4

<212> PRT

<213> Artificial sequence

<220>

<223> Synthetic construct

<220>
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<222> (2)..(2)
<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
<221> MISC_FEATURE
<222> (3)..(3)
<223> Xaa is Atc, i.e. 2-aminotetralin-2-carboxylic acid

<400> 35

Tyr Xaa Xaa Phe
1

<210> 36
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<213> Artificial sequence

<220>
<223> Synthetic construct

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<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
<221> MOD_RES
<222> (4)..(4)
<223> Phe is Phe(pNO2), i.e. 4-nitrophenylalanine

<400> 36

Tyr Xaa Phe Phe
1

<210> 37
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<220>
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<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
<221> MOD_RES

<222> (4)..(4)
<223> Phe is Phe(pNO2), i.e. 4-nitrophenylalanine

<400> 37

Tyr Xaa Trp Phe
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<220>
<223> Synthetic construct

<220>
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<222> (2)..(2)
<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
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<222> (4)..(4)
<223> AMIDATION

<400> 38

Tyr Xaa Phe Trp
1

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<223> Synthetic construct

<220>
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<222> (2)..(2)
<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
<221> MOD_RES
<222> (7)..(7)
<223> AMIDATION

<400> 39

Tyr Xaa Phe Phe Val Val Gly
1 5

<210> 40
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<220>
<223> Synthetic construct

<220>
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<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
<221> MOD_RES
<222> (7)..(7)
<223> AMIDATION

<400> 40

Tyr Xaa Phe Phe Tyr Pro Ser
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<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
<221> MOD_RES
<222> (7)..(7)
<223> AMIDATION

<400> 41

Tyr Xaa Trp Phe Tyr Pro Ser
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<210> 42
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<213> Artificial sequence

<220>
<223> Synthetic construct

<220>
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<222> (2)..(2)
<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
<221> MOD_RES
<222> (4)..(4)
<223> Phe is Phe(pNO2), i.e. 4-nitrophenylalanine

<220>
<221> MOD_RES
<222> (7)..(7)
<223> AMIDATION

<400> 42

Tyr Xaa Trp Phe Tyr Pro Ser
1 5

<210> 43
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<220>
<223> Synthetic construct

<220>
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<223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
<221> MOD_RES
<222> (6)..(6)
<223> Nle

<220>
<221> MOD_RES
<222> (7)..(7)
<223> AMIDATION

<400> 43

Tyr Xaa Phe Phe Leu Leu Asp
1 5

<210> 44
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<212> PRT
<213> Artificial sequence

<220>
 <223> Synthetic construct

 <220>
 <221> MISC_FEATURE
 <222> (2)..(2)
 <223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

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Tyr Xaa Phe
 1

<210> 45
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<220>
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<220>
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 <222> (2)..(2)
 <223> Xaa is Tic, i.e. 1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
 <221> MOD_RES
 <222> (3)..(3)
 <223> AMIDATION

<400> 45

Tyr Xaa Phe
 1

<210> 46
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<220>
 <223> Synthetic construct

<220>
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 <222> (2)..(2)
 <223> Xaa is Tic-psi-[CH2-], i.e. 3-methyl-1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>
 <221> MOD_RES
 <222> (2)..(3)
 <223> nonpeptidyl bond

<400> 46

Tyr Xaa Phe

1

<210> 47

<211> 4

<212> PRT

<213> Artificial sequence

<220>

<223> Synthetic construct

<220>

<221> MISC_FEATURE

<222> (2)..(2)

<223> Xaa is Tic-psi-[CH2-], i.e. 3-methyl-1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid

<220>

<221> MOD_RES

<222> (2)..(3)

<223> nonpeptidyl bond

<400> 47

Tyr Xaa Phe Phe

1

<210> 48

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<213> Artificial sequence

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<400> 48

Tyr Gly Gly Phe Met

1

5

<210> 49

<211> 6

<212> PRT

<213> Artificial sequence

<220>

<223> Synthetic construct

<400> 49

Tyr Gly Gly Phe Met Lys

1

5

<210> 50
<211> 6
<212> PRT
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<220>
<223> Synthetic construct

<220>
<221> MOD_RES
<222> (1)..(1)
<223> NH2 of Tyr is blocked by butyloxycarbonyl group

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Tyr Gly Gly Phe Leu Lys
1 5

<210> 51
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<220>
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<220>
<221> MOD_RES
<222> (1)..(1)
<223> NH2 of Tyr is blocked by butyloxycarbonyl group

<400> 51

Tyr Gly Gly Phe Leu Lys
1 5

<210> 52
<211> 6
<212> PRT
<213> Artificial sequence

<220>
<223> Synthetic construct

<220>
<221> MOD_RES
<222> (1)..(1)
<223> NH2 of Tyr is blocked by butyloxycarbonyl group

<220>
<221> MOD_RES
<222> (6)..(6)
<223> Polymer connected to epsilon-amino group

<400> 52

Tyr Gly Gly Phe Leu Lys
1 5

<210> 53

<211> 2

<212> PRT

<213> Artificial sequence

<220>

<223> Synthetic construct

<220>

<221> MOD_RES

<222> (1)..(1)

<223> Tyr has at its N-terminus a Me-x-H-y-N group, wherein x is 0, 1, or 2; and y is 0, 1, or 2, with the proviso that x and y is never greater than 2

<220>

<221> MOD_RES

<222> (1)..(2)

<223> The amine between the first Tyr and the second Tyr is methylated, wherein z is 0 or 1

<220>

<221> MOD_RES

<222> (2)..(2)

<223> AMIDATION

<220>

<221> MOD_RES

<222> (2)..(2)

<223> The amidated C-terminal is benzylated (NHBzl)

<400> 53

Tyr Tyr
1